

SUPPLEMENTAL COMBINED DECLARATION AND POWER OF ATTORNEY

As a below named inventor, I hereby declare that:

My residence, post office address and citizenship are as stated below next to my name; and

I verily believe that I am the original, first and sole inventor (if only one name is listed below) or an original, first and joint inventor (if plural names are listed below) of the subject matter which is claimed and for which a patent is sought on the invention entitled: **'Oligonucleotides Containing 2'-O-Modified Purines,** the specification of which:

() is attached hereto.

(XX) was filed on **October 27, 1992**, as Application Serial No. **07/967,267** and was amended on **January 3, 1994** and **July 7, 1994**.

I hereby state that I have reviewed and understand the contents of the above identified specification, including the claims, as amended by any amendment referred to above.

I acknowledge the duty to disclose to the U.S. Patent and Trademark Office all information known to be material to the patentability of this application in accordance with 37 CFR § 1.56.

I hereby claim foreign priority benefits under 35 U.S.C. § 119 of any foreign application(s) for patent or inventor's certificate listed below and have also identified below any foreign application for patent or inventor's certificate having a filing date before that of any application on which priority is claimed:

Country	Number	Date Filed	Priority Claimed
<u>PCT</u>	<u>US91/00243</u>	<u>January 1, 1991</u>	<u>Yes.</u>

I hereby claim the benefit under 35 U.S.C. § 120 of any United States application(s) listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States application in the manner provided by the first paragraph of 35 U.S.C. § 112, I acknowledge the duty to disclose to the U.S. Patent and Trademark Office all information known to be material to patentability as defined in 37 CFR § 1.56 which became available between the filing date of the prior application and the national or PCT international filing date of this application:

Application Serial No.	Filing Date	Status (patented, pending)
<u>07/918,362</u>	<u>July 23, 1992</u>	<u>pending</u>
<u>07/463,358</u>	<u>January 11, 1990</u>	<u>abandoned</u>
<u>07/566,977</u>	<u>August 13, 1990</u>	<u>abandoned</u>

I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and to transact all business in the Patent and Trademark Office connected therewith: **John W. Caldwell, Esquire** and **Joseph Lucci, Registration Nos. 28,937 and 33,307** of the firm of **WOODCOCK WASHBURN KURTZ MACKIEWICZ & NORRIS**, One Liberty Place - 46th Floor, Philadelphia, Pennsylvania 19103, and **Herb Boswell and Laurel Bernstein, Registration Nos. 27,311 and 37,280**, of **ISIS Pharmaceuticals**, 2280 Faraday Avenue, Carlsbad, California 92008.

Address all telephone calls and correspondence to:

Joseph Lucci, Esquire
WOODCOCK WASHBURN KURTZ MACKIEWICZ & NORRIS
One Liberty Place - 46th Floor
Philadelphia, PA 19103
Telephone No. 215-568-3100.

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the

United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

1	Full Name Phillip Dan Cook	Inventor's Signature <i>Phillip Dan Cook</i>	Date 3/06/75
	Residence San Marcos, CA 92069	Citizenship U.S.A.	
	Post Office Address 641 Avilar Court		
2	Full Name Daniel Peter Claude McGee	Inventor's Signature <i>Daniel Peter Claude McGee</i>	Date 7 MARCH '95
	Residence Boulder, CO 80301	Citizenship U.S.A. - CANADIAN <i>DPM</i>	
	Post Office Address 7715 Devonshire Way		
3	Full Name Charles John Guinossos	Inventor's Signature <i>Charles John Guinossos</i>	Date 6 MAR 95
	Residence Vista, CA 92083	Citizenship U.S.A.	
	Post Office Address 313 Encino Drive		
4	Full Name	Inventor's Signature	Date
	Residence	Citizenship	
	Post Office Address		
5	Full Name	Inventor's Signature	Date
	Residence	Citizenship	
	Post Office Address		

COMBINED DECLARATION AND POWER OF ATTORNEY

As a below named inventor, I hereby declare that:

My residence, post office address and citizenship are as stated below next to my name; and

I verily believe that I am the original, first and sole inventor (if only one name is listed below) or an original, first and joint inventor (if plural names are listed below) of the subject matter which is claimed and for which a patent is sought on the invention entitled: NOVEL 2' O-ALKYL GUANOSINES the specification of which:

(xx) is attached hereto.

() was filed on _____ as Application Serial No. _____
and was amended on _____ (if applicable).

I hereby state that I have reviewed and understand the contents of the above identified specification, including the claims, as amended by any amendment referred to above.

I acknowledge the duty to disclose to the U.S. Patent and Trademark Office all information known to be material to the patentability of this application in accordance with 37 CFR § 1.56.

I hereby claim foreign priority benefits under 35 U.S.C. § 119 of any foreign application(s) for patent or inventor's certificate listed below and have also identified below any foreign application for patent or inventor's certificate having a filing date before that of any application on which priority is claimed:

Country	Number	Date Filed	Priority Claimed
_____	_____	_____	_____
_____	_____	_____	_____
_____	_____	_____	_____

I hereby claim the benefit under 35 U.S.C. § 120 of any United States application(s) listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States application in the manner provided by the first paragraph of 35 U.S.C. § 112, I acknowledge the duty to disclose to the U.S. Patent and Trademark Office all information known to be material to patentability as defined in 37 CFR § 1.56 which became available between the filing date of the prior application and the national or PCT international filing date of this application:

Application Serial No.	Filing Date	Status (patented, pending)
<u>918,362</u>	<u>July 23, 1992</u>	<u>pending</u>

I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and to transact all business in the Patent and Trademark Office connected therewith: John W. Caldwell and Rebecca R. Gaumond, Registration Nos. 28,937 and 35,152 of the firm of WOODCOCK WASHBURN KURTZ MACKIEWICZ & NORRIS, One Liberty Place - 46th Floor, Philadelphia, Pennsylvania 19103, and Herb Boswell, Registration No. 27,311, of ISIS Pharmaceuticals, 2280 Faraday Avenue, Carlsbad, California 92008.

Address all telephone calls and correspondence to:

Rebecca R. Gaumond, Esq.
WOODCOCK WASHBURN KURTZ MACKIEWICZ & NORRIS
One Liberty Place - 46th Floor
Philadelphia, PA 19103
Telephone No. 215-568-3100.

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the

United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

	Full Name Phillip Dan Cook	Inventor's Signature <i>Phillip Dan Cook</i>	Date 10/13/61
1	Residence 7340 Bolero Street Carlsbad, CA 92009	Citizenship	US
	Post Office Address Same as above		
	Full Name Daniel Peter Claude McGee	Inventor's Signature <i>Daniel Peter Claude McGee</i>	Date 23 Oct 1962
2	Residence 295 Chinquapin Avenue Carlsbad, CA 92008	Citizenship	Canadian
	Post Office Address Same as above		
	Full Name Charles John Guinosso	Inventor's Signature <i>Charles John Guinosso</i>	Date 23 Oct 1962
3	Residence 3399 Calle de Sur Carlsbad, CA 92009	Citizenship	US
	Post Office Address Same as above		
	Full Name	Inventor's Signature	Date
4	Residence	Citizenship	
	Post Office Address		
	Full Name	Inventor's Signature	Date
5	Residence	Citizenship	
	Post Office Address		